

**IN THE CLAIMS:**

A listing of the claims, in accordance with 37 CFR §1.121, is provided. The listing of claims replaces all prior such listings of claims. Claims 8, 14 and 58 are amended herein.

Claims 73 and 74 are added herein.

Claims 1-7 (Cancelled)

8. (Currently Amended) A high-throughput method of assigning a function ~~corresponding to a phenotype and~~ associated with a product coded for by a ~~nucleotide sequence of a sample nucleic acid sequence in a target nucleic acid molecule~~, said method comprising:

a) without any intervening bacterial cloning steps and without any conformational modeling of mRNA transcribed from the target nucleic acid molecule that comprises the sample nucleic acid sequence, delivering into and amplifying and expressing one or more a plurality of members of an oligonucleotide family as individual transcription products in a plurality of recombinant non-bacterial host cells comprising [[a]] the target nucleic acid molecule that comprises the nucleotide sequence of the sample nucleic acid sequence, whereby the method is high-throughput, wherein:

the members of the oligonucleotide family comprise a plurality of nucleic acids each encoding a transcription product comprising a sequence that is complementary to a sequence contained in the mRNA transcribed from the target nucleic acid molecule that comprises the sample nucleic acid sequence;

the coding sequences for each individual transcription product encodes an antisense nucleic acid that, when expressed as RNA, binds to the mRNA transcribed from the target nucleic acid molecule that comprises the ~~nucleotide sequence of the sample nucleic acid sequence~~; and

expression of one or more of the individual transcription products inhibits production of a product of the mRNA; and

b) in the resulting host cells, analyzing changes in ~~the phenotype to thereby assign a function associated with the product encoded by the nucleotide sequence of the sample nucleic acid sequence in the target nucleic acid molecule~~.

9. (Original) The method according to Claim 8, wherein said function is a physiological function.

10. (Original) The method according to Claim 8, wherein said function is enzyme activity.

11. (Original) The method according to Claim 8, wherein said function is protein synthesis.

12. (Original) The method according to Claim 8, wherein said function is expression of a biological factor.

13. (Original) The method according to Claim 8, wherein said function is a regulatory effector function.

14. (Currently Amended) The method according to Claim 8, wherein said ~~function phenotypic change is changed monitored~~ directly.

Claims 15-57 (Cancelled)

58. (Currently Amended) The method of claim 8, wherein the ~~one or more~~ plurality of members of the oligonucleotide family are introduced into expression vectors, which are introduced into the host cells, wherein the expression vectors comprise:

double-stranded DNA, comprising:

a sense strand and an antisense strand, wherein the sense strand codes for an antisense strand that, when expressed as RNA, binds to an mRNA sequence transcribed from the target nucleic acid sequence so that expression of a product from the target nucleic acid is inhibited; and

means for determining directionality of expression, wherein the product is associated with at least one phenotypic property of a host cell containing the mRNA sequence; and wherein the expression vector is for expression in non-bacterial host cells.

59. (Previously Presented) The method of claim 58, wherein the RNA comprises:

a catalytic domain that cleaves an mRNA sequence transcribed from the target nucleic acid; and

binding sequences flanking the catalytic domain for binding the RNA to the mRNA, and/or wherein the means for determining directionality of expression comprises a different non blunt-ended restriction enzyme site at each end of said double-stranded DNA.

60. (Original) The method of claim 59, wherein the double-stranded DNA is formed by contacting a first oligonucleotide with a complementary second oligonucleotide, and/or wherein the non blunt-ended restriction enzyme site is complementary to an end of the expression vector.

61. (Original) The method of claim 59, wherein said expression vector is formed by: (a) contacting a double-stranded oligonucleotide with an expression vector; or (b) by contacting a single-stranded oligonucleotide with said expression vector; or (c) contacting a triple-stranded oligonucleotide with an expression vector.

62. (Previously Presented) The method of claim 58, wherein the expression vector is a plasmid or a virus for expression in non-bacterial host cells.

63. (Original) The method of claim 62, wherein the virus is a retrovirus or an adeno-associated virus.

64. (Previously Presented) The method of claim 58, wherein the expression vector is transfected directly into mammalian cells.

65. (Previously Presented) The method of claim 8, wherein the sample nucleic acid is genomic DNA, cDNA, an expressed sequence tag (EST) or RNA.

66. (Previously Presented) The method of claim 8, wherein the family contains between 3 and 20 members.

67. (Previously Presented) The method of claim 8, wherein each member of the family is designed to inhibit the production of a product of the target nucleic acid molecule.

68. (Previously Presented) The method of claim 8 that is performed in a high throughput format, whereby all members of a family are assessed in a single experiment.

69. (Previously Presented) The method of claim 8 that is performed in a high throughput format, whereby a plurality of different target nucleic acid molecules and/or sample nucleotide sequences are assessed.

70. (Original) The method of claim 59, wherein the expression vector is a plasmid or a virus for expression in non-bacterial host cells.

71. (Original) The method of claim 60, wherein the expression vector is a plasmid or a virus for expression in non-bacterial host cells.

Applicant : James Keck, et. al  
Serial No. : 09/601,997  
Filed : December 15, 2000  
Amendment in Response to Office Action

Attorney's Docket No.: 17111-007US1 (2307US)

72. (Original) The method of claim 61, wherein the expression vector is a plasmid or a virus for expression in non-bacterial host cells.

73. (New) The method of claim 8, wherein the oligonucleotide family is a ribozyme family.

74. (New) The method of claim 73, wherein the sequence contained in the transcription product encoded by the oligonucleotide family member that is complementary to a sequence contained in the mRNA transcribed from the target nucleic acid molecule flanks a catalytic domain sequence based on a consensus cleavage site identified in the mRNA transcribed from the target nucleic acid molecule.